

Allergy School on Hymenoptera Venom Allergy 2013

Insect stings and bites - not only hymenoptera



Prof.

Peter Schmid-Grendelmeier

Allergy Unit, Dept. Of Dermatology
University Hospital
Zuerich, Switzerland



Insect bites and stings

Most common allergenes

Clinical features

Cross-reactivity

Management



Insect bites and stings

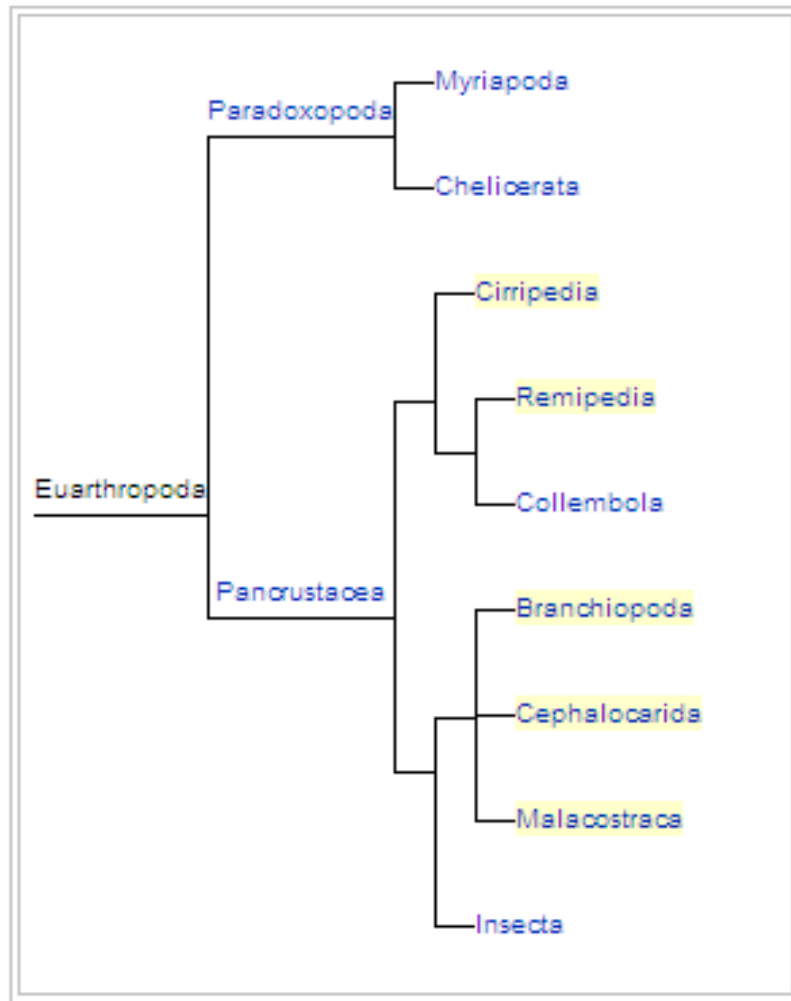
Pain /Local reaction

Allergy

Infections / Transmission



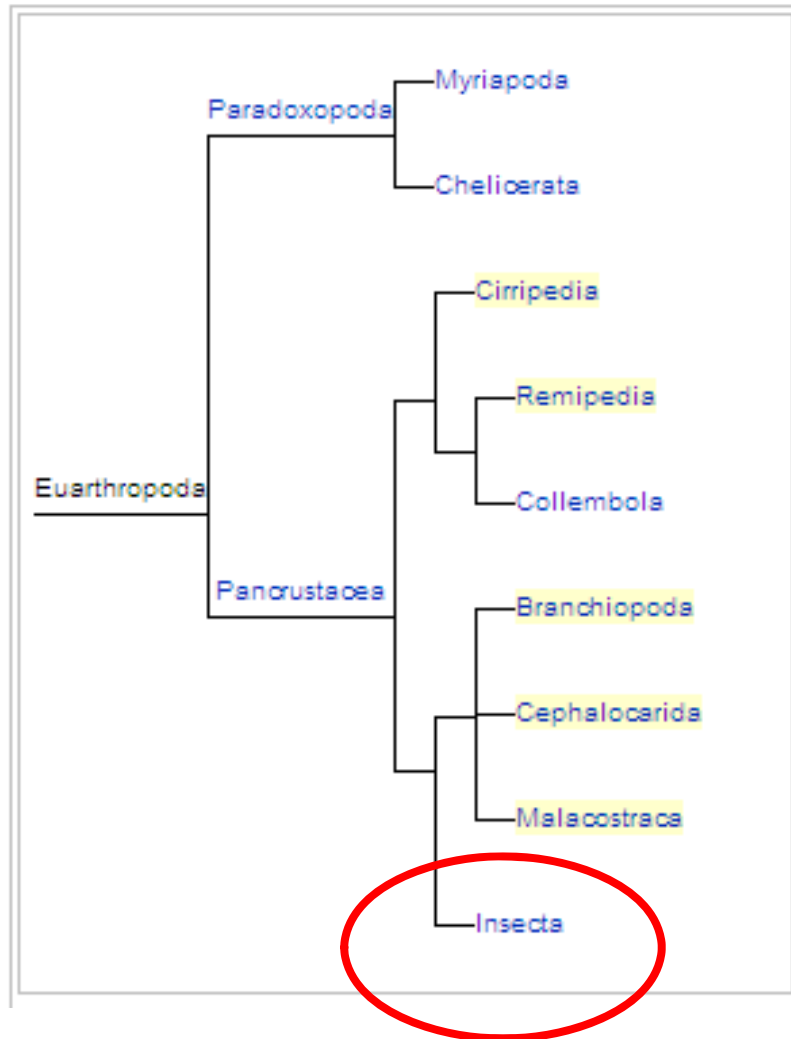
Arthropods



Arthropods



Spiders, Mites, Ticks, Scorpions



Centipeds, Millipeds



Crustaceae

Shrimps, Crabs, Lobster, Crabfish, Woodlice



Nocturnal asthma

Feeling of stings/bites







Bites of the European pigeon tick (*Argas reflexus*): Risk of IgE-mediated sensitizations and anaphylactic reactions

Jörg Kleine-Tebbe, MD,^a Anja Heinatz, MD,^a Inken Gräser, MD,^a Hans Dautel, PhD,^b
Gitte Nordskov Hansen, BSc,^c Sabine Kespohl, PhD,^d Hans-Peter Rihs, PhD,^d
Monika Raulf-Heimsoth, PhD,^d Günther Vater, PhD,^e Manfred Rytter, MD,^a and
Uwe-Fritjof Haustein, MD^a *Leipzig, Berlin, and Bochum, Germany, and Hørsholm, Denmark*



Allergy to pigeon tick (*Argas reflexus*)

TABLE I. LRs and SRs after a bite of the pigeon tick *Argas reflexus* in 148 subjects (37 atopics, 24%, and 111 nonatopics)

Symptoms	Frequency	
	n	%
Local	146	99
Redness	142	96
Local itching	124	84
Inflammatory node	93	63
Lymphatic secretion	36	24
Wheal	25	17
Pruritic secretion	20	14
Lymphangitis	12	8
Lymph node swelling	10	7
Vesicle	10	7
Systemic	12	8
Urticaria	11	7
Angioedema	6	4
Vascular dysregulation	2	1
Dyspnea	7	5
Unconsciousness	4	3
Gastrointestinal symptoms	3	2
Rush with pruritus	3	2

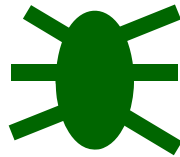
Mostly local reaction

About 8% systemic



Arthropods

Insects



6 legs

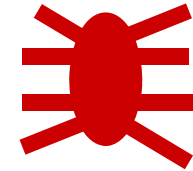
Flies

Mosquitos etc

Hymenoptera

Bee, Wasp, Ants

Arachnoidea



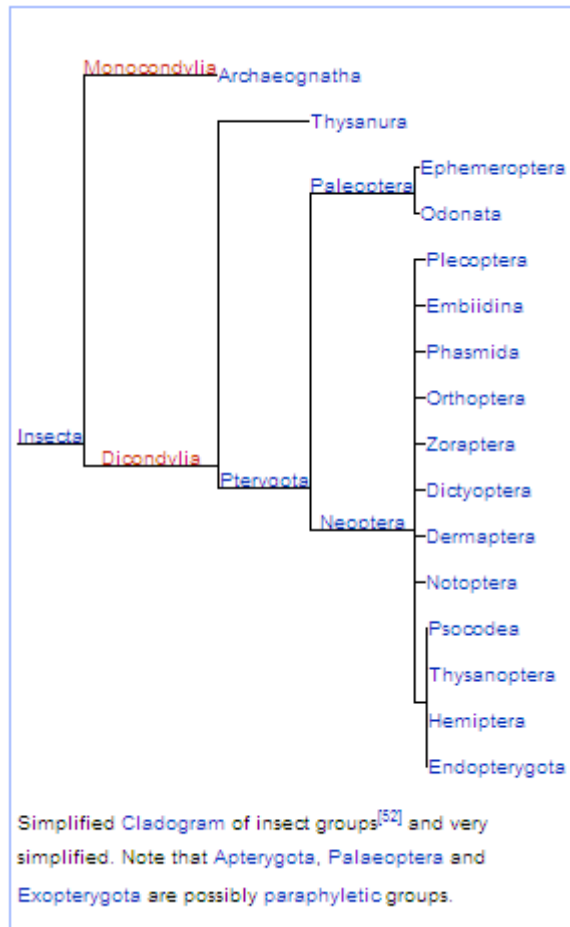
8 legs

Spiders

Ticks

Scorpions

Insects

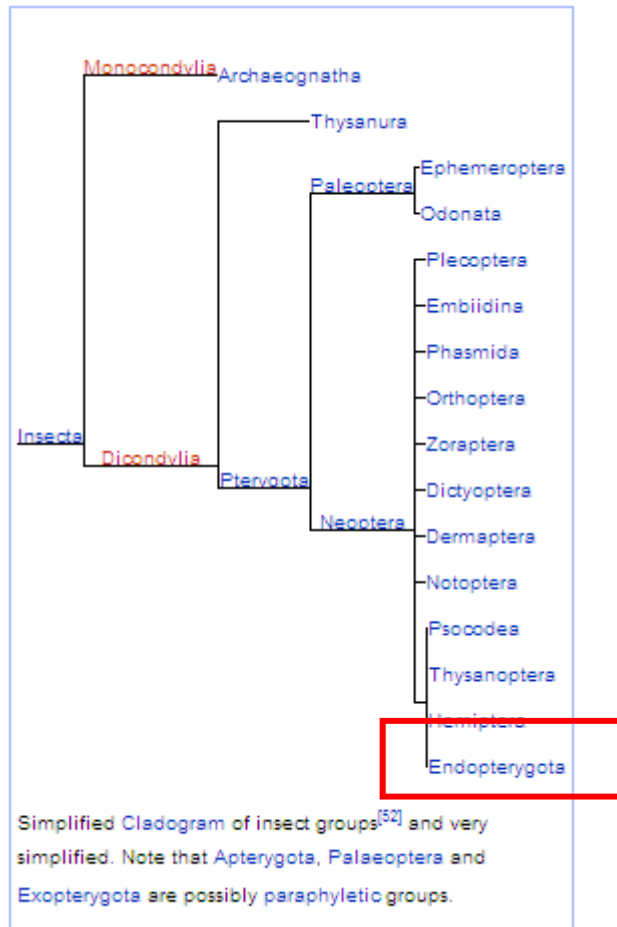


6- 7 Mio Species

About 1 Mio known

➤ 170'000 butterflies

> 350'000 beetles



- Hymenoptera (ants, bees, etc.)
- Coleoptera (beetles)
- Strepsiptera (twisted-winged parasites)

K.U, m, 29-j.

Camping holidays in Florida, USA

Feeling of „insect bites“

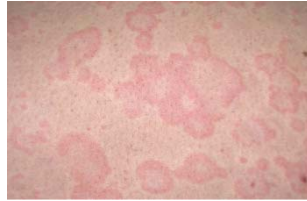
Suddenly Urticaria, Sweating, Dyspnoe

Hospital admission



Severity of Anaphylaxis (H.L. Muller)

Grade I



Urticaria

Grade II



Angio-Edema, Nausea,
Vomiting,
Diarrhoe

Grade III



Dyspnoea, Cough, Stridor
Dysphagia, Weakness,
Confusion

Grade IV



Hypotension,
Kollaps, Incontinence
Loss of consciousness,
Cyanosis



K.U, m, 29-j.

Allergologic work up



	Bee	Wasp	Imported Red Fire ant
Spec IgE in Serum			
Class	0	2	3
kU/l		0.98	1.89
Intradermal skin test		pos. at 1.0 ug/ml	
Normal		1.0	

Red imported fire ant



Red imported fire ant *Solenopsis invicta*

Originally from **Brazil**



Mato Grosso



Red imported fire ant *Solenopsis invicta*

Originally from **Brazil**

Mato Grosso



Red imported fire ant *Solenopsis invicta*

Originally from **Brazil**

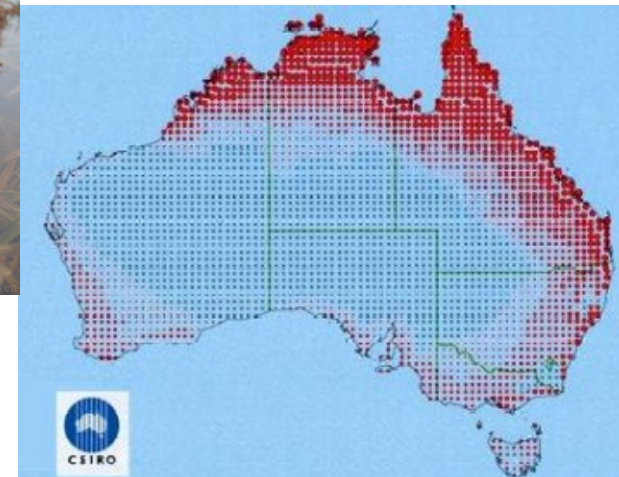
Around 1940 **Southern USA**



Red imported fire ant *Solenopsis invicta*

Originally from **Brazil**

Around 1940 **Southern USA / Australia**



Red imported fire ant are not the largest ants

Comparison of ant sizes



Fire ant
Solenopsis invicta



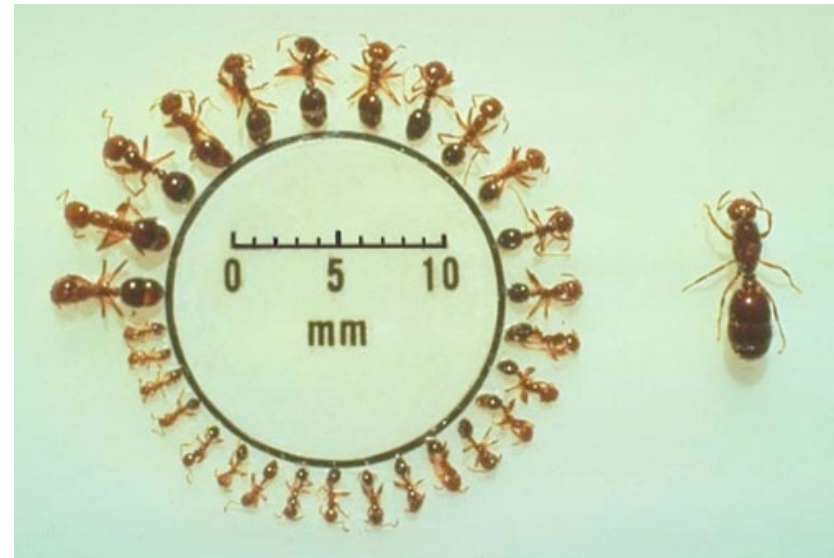
Black ant
Iridomyrmex 'rufoniger'



Sugar ant
Camponotus sp.



Bull ant
Myrmecia sp.



2- 6 mm

Red imported fire ant

but

- spread fast
- are invasive and extinct other ants
- are aggressive

Meet the Fire Ant: An Introduction to an Exotic Pest

Megan Bame

Pest animals

The term 'pest animal' refers to any exotic animal, which causes a detrimental impact on the environment, industry or community activities.

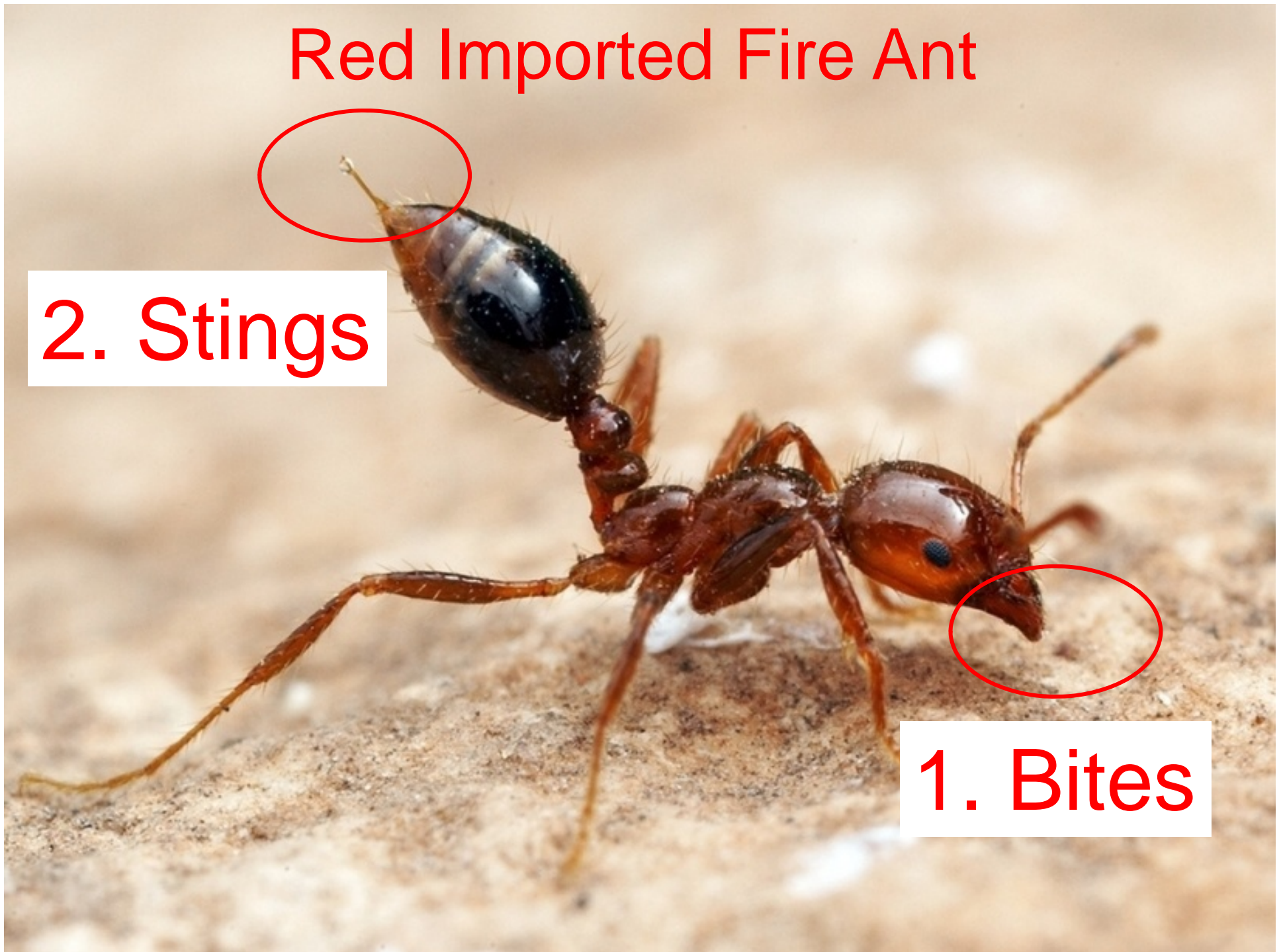
Pest species may come from any animal group including mammals, birds, fish, reptiles, amphibians, crustaceans, molluscs and insects.



Red Imported Fire Ant

2. Stings

1. Bites



Red Imported Fire Ant

2. Stings

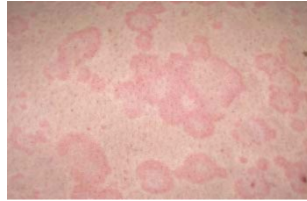


1. Bites



Allergy to RFA (H.L. Muller)

Grade I



Urticaria



Grade II



Angio-Edema, Nausea,
Vomiting,
Diarrhœe

Grade III

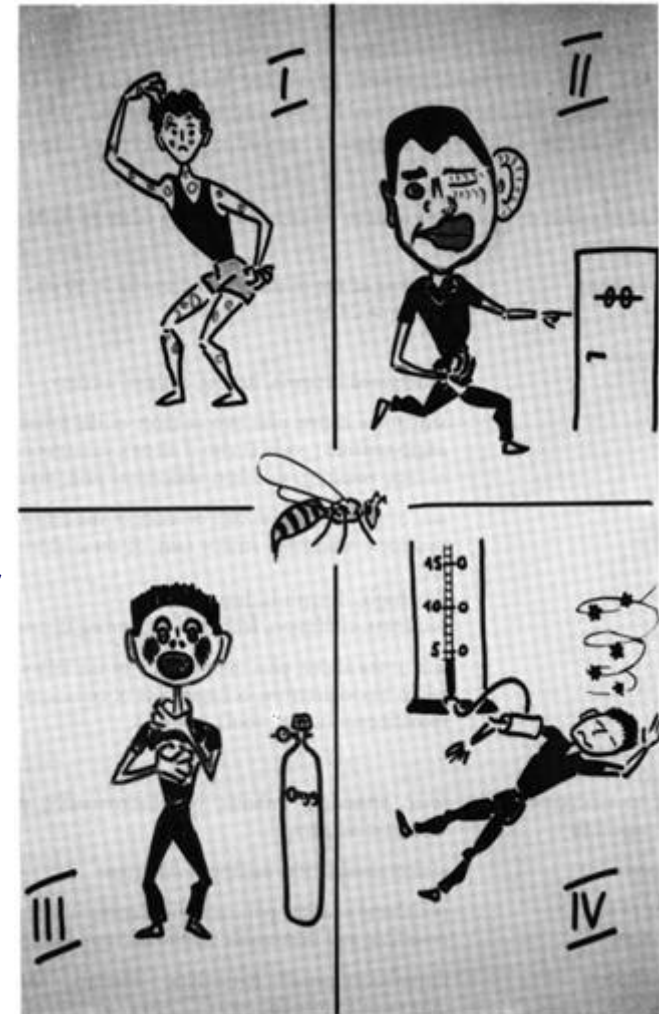


Dyspnoea, Cough, Stridor
Dysphagia, Weakness,
Confusion

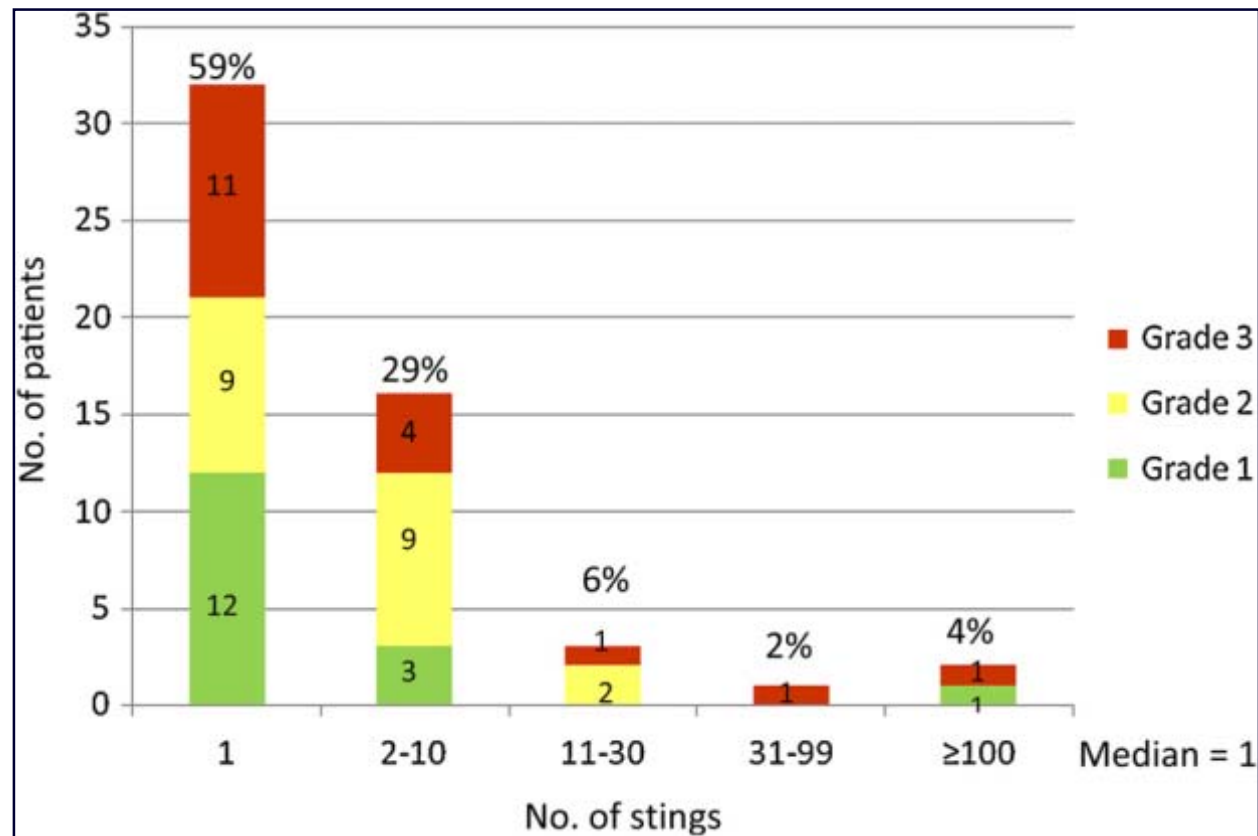
Grade IV



Hypotension,
Kollaps, Incontinence
Loss of consciousness,
Cyanosis



Allergic reactions to RFA often occur at early stings



Allergens of RFA

Sol i 1

Sol i 2

Sol i 3

Sol i 4

Hoffman DR et al J Allergy Clin Immunol 1988;82:828-34
Stafford T. Ann Allergy Asthma Immunol 1996;77:87-99.

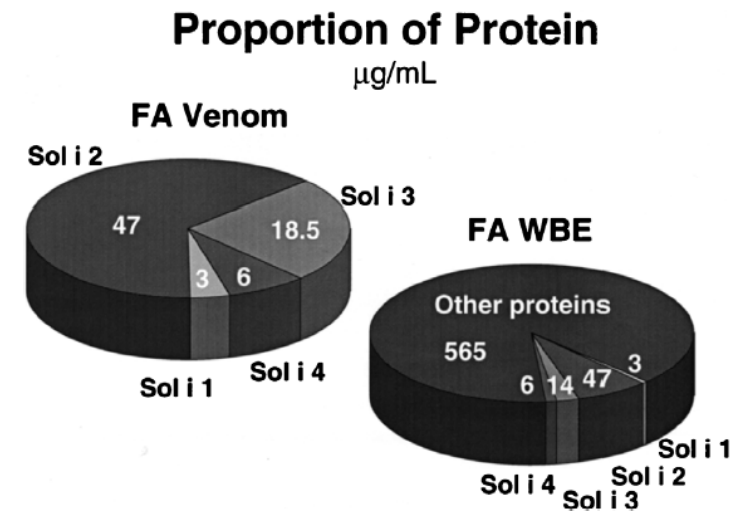


Figure 4. *Left:* Concentration Sol i allergens in fire ant venom. *Right:* Sol i allergen concentration compared to relatively large amount of extraneous body protein in fire ant whole body extract.

Allergens of RFA

Sol i 1 Phospholipase
cross-reacts with
Wasps

Sol i 2 specific for Solenopsis

Sol i 3

Sol i 4 specific for Solenopsis

Hoffman DR et al J Allergy Clin Immunol 1988;82:828-34

Stafford T. Ann Allergy Asthma Immunol 1996;77:87-99.

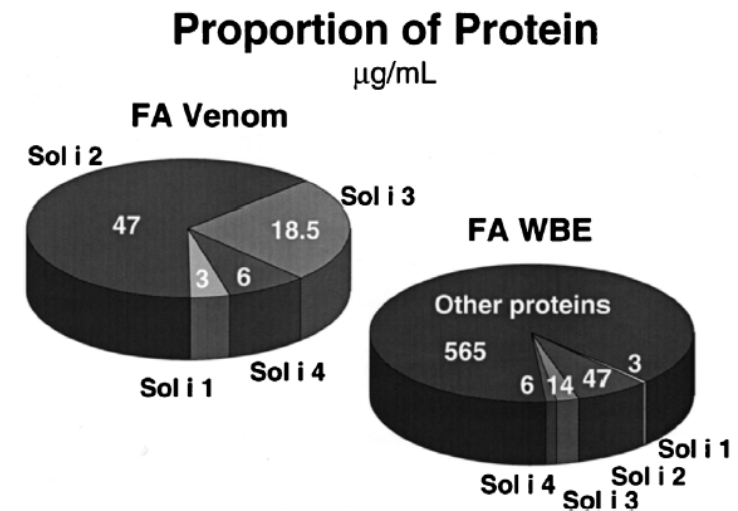


Figure 4. *Left:* Concentration Sol i allergens in fire ant venom. *Right:* Sol i allergen concentration compared to relatively large amount of extraneous body protein in fire ant whole body extract.

RFA

Diagnosis of Allergy

Specific IgE in Serum

ImmunoCAP Code: i70



Source material: Whole body extract

Skin tests



Prick test (Commercial extracts)

Intradermal tests (Commercial extracts)

Source material: Whole body extract

Solley Med J Aust 2002; 176: 518

La Shell M et al J Allergy Clin Immunol 2010;125:1294-9

RFA

Treatment

Grad I-II

Emergency kit

Grad III- IV

Spec. Immunotherapy

Source material: Whole body extract



Solley Med J Aust 2002; 176: 518

La Shell M et al J Allergy Clin Immunol 2010;125:1294-9

Imported fire ant field reaction and immunotherapy safety characteristics: The IFACS study

Mark S. La Shell, MD,^a Christopher W. Calabria, MD,^b and James M. Quinn, MD^b *Travis Air Force Base, Calif, and Lackland Air Force Base, San Antonio, Tex*

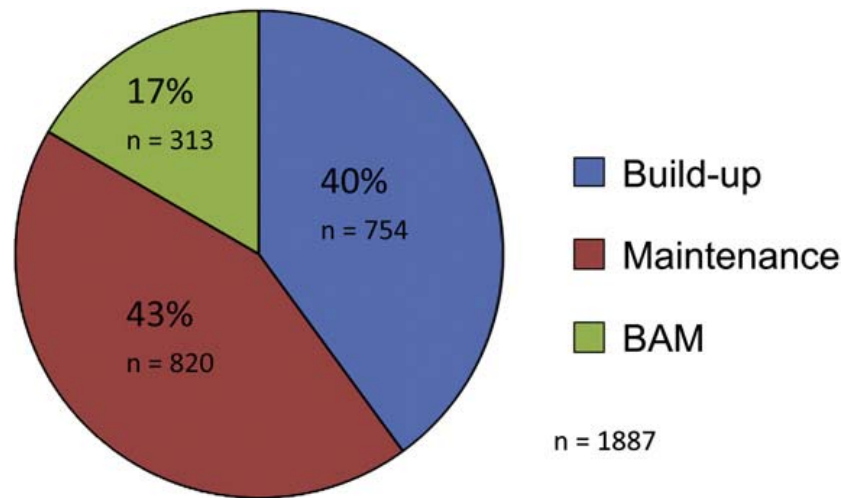


FIG 2. Total injections by phase.

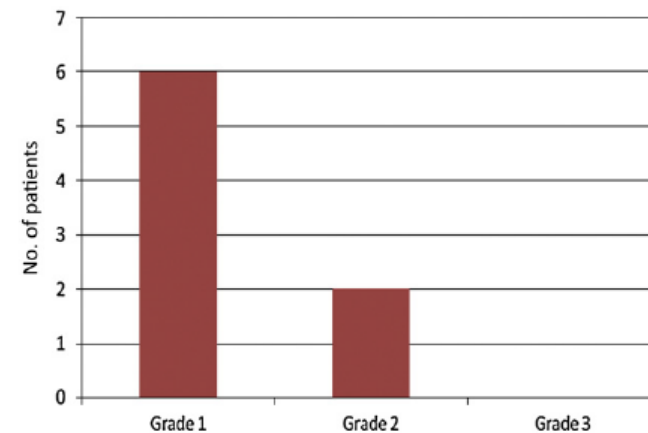














FIG 3. Severity of SRs to IFA SCIT. Seven of 77 patients experienced SRs to IFA SCIT. There were no emergency department visits, hospitalizations, or fatalities. One patient experienced more than 1 SR, and she had a total of 2 SRs, each grade 1. In 2 different patients 2 SRs were delayed (one presented at 35 minutes and the other at 60 minutes).

Imported fire ant field reaction and immunotherapy safety characteristics: The IFACS study

Mark S. La Shell, MD,^a Christopher W. Calabria, MD,^b and James M. Quinn, MD^b *Travis Air Force Base, Calif, and Lackland Air Force Base, San Antonio, Tex*

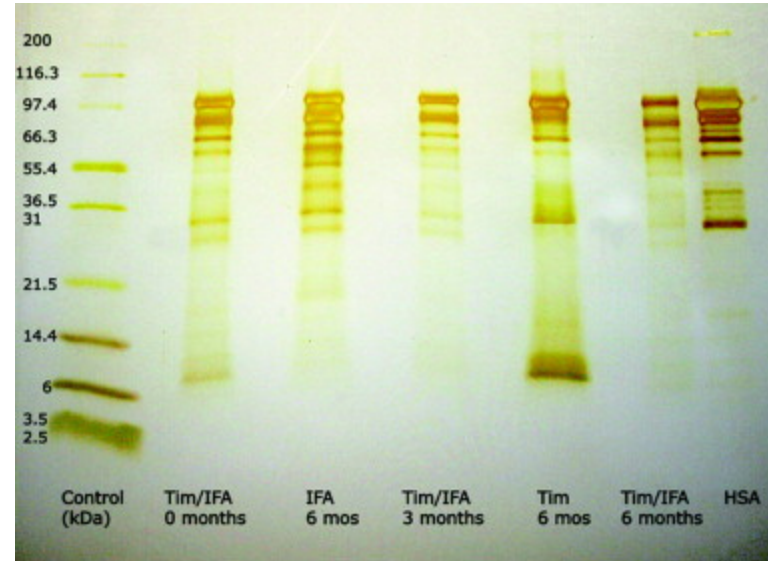
TABLE III. Risk factors for SRs to IFA SCIT per injection

	SR to IFA SCIT	No SR to IFA SCIT		OR	95% CI
LLR	2 (25%)	18 (1.0%)		34.4*	6.51-182
Asthma	2 (25.0%)	523 (27.8%)		0.87	0.18-4.32
Sex (female)	6 (75%)	1,071 (57.0%)		2.29	0.46-11.4
Age (>50 y)	0 (0%)	666 (35.4%)		0.11	0.01-1.98
ACE inhibitor	0 (0%)	173 (9.21%)		0.62	0.04-10.8
Field reaction†	3 (37.5%)	608 (36.4%)		1.04	0.25-4.39
Maintenance‡	3 (37.5%)	817 (43%)		0.78	0.19-3.27
Build-up and BAM‡	5 (62.5%)	1,062 (59.7%)		1.28	0.31-5.38
Build-up‡	2 (25.0%)	752 (42%)		0.5	0.10-2.48
BAM‡	3 (37.5%)	310 (18%)		3.04	0.72-12.8
SR to ST	3 (37.5%)	175 (11.1%)		4.75*	1.13-20.0
Concentration of ST§	8 (100%)	0 (0%)		4.33	0.25-75.7

0.01 0.2 1 10 100 1000

SCIT with RFA WHBE is safe,
but increased risk in patients with LLR
with SR to Skin tests

Compatibility of imported fire ant whole body extract with cat, ragweed, *Dermatophagoides pteronyssinus*, and timothy grass allergens



Conclusions

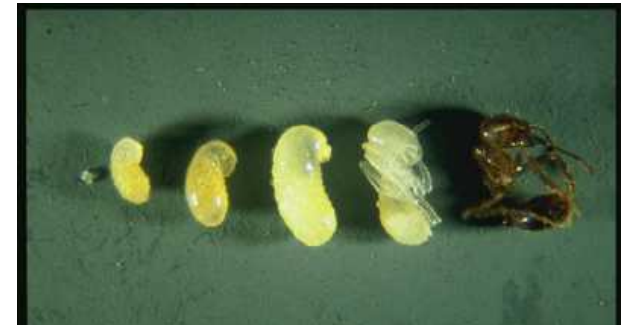
Imported fire ant whole body extract combined with timothy grass resulted in significant and rapid timothy protein degradation.

Imported fire ant whole body extract mixed with cat, ragweed, or *D pteronyssinus* revealed aeroallergen stability, yielding the possibility of combining these extracts in a single immunotherapy injection.

Compatibilities of IFA WBE with other common aeroallergens remain undetermined and thus **are not recommended for single-injection immunotherapy formulations.**

Rans Tet al Ann Allergy Asthma Immunol. 2009;102:57–61.

Red imported fire ant *Solenopsis invicta*



J Invest Allergol Clin Immunol 2006

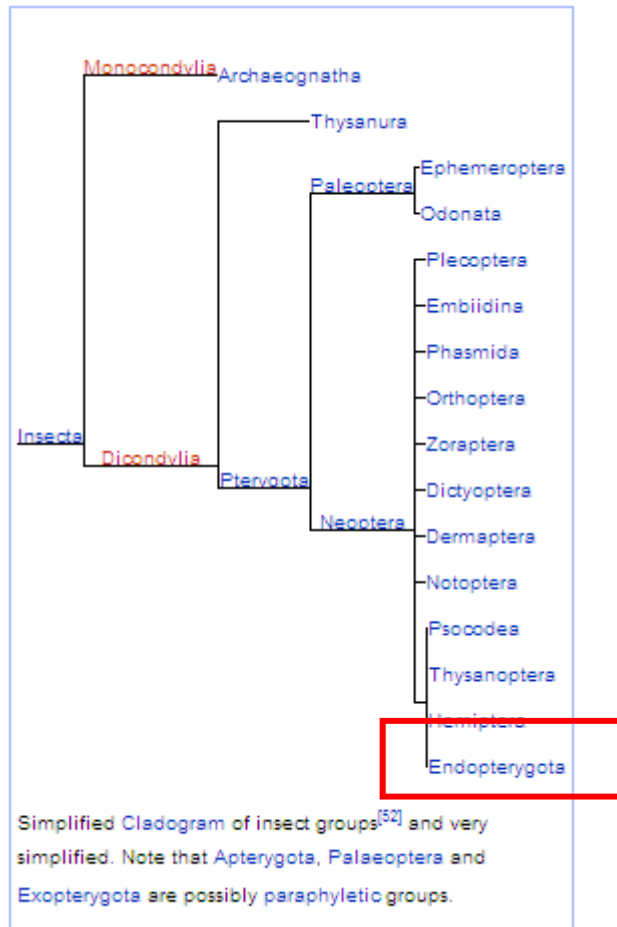


CASE REPORT

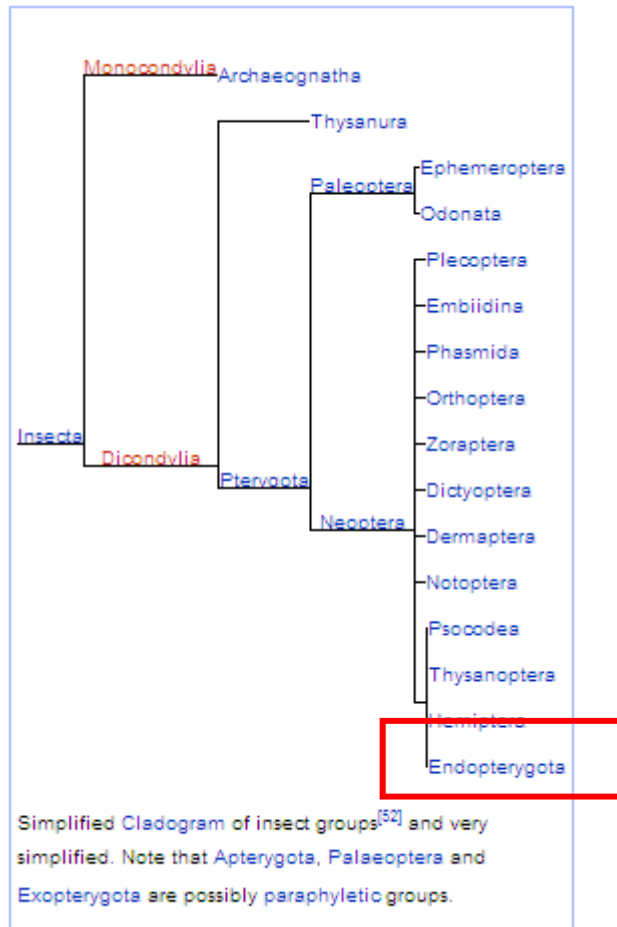
Anaphylaxis Caused by Imported Red Fire Ant Stings in Málaga, Spain

S Fernández-Meléndez,¹ A Miranda,¹ JJ García-González,¹ D Barber,²
M Lombardero²

Due to work with tropical wood
imported from Brazil to Spain

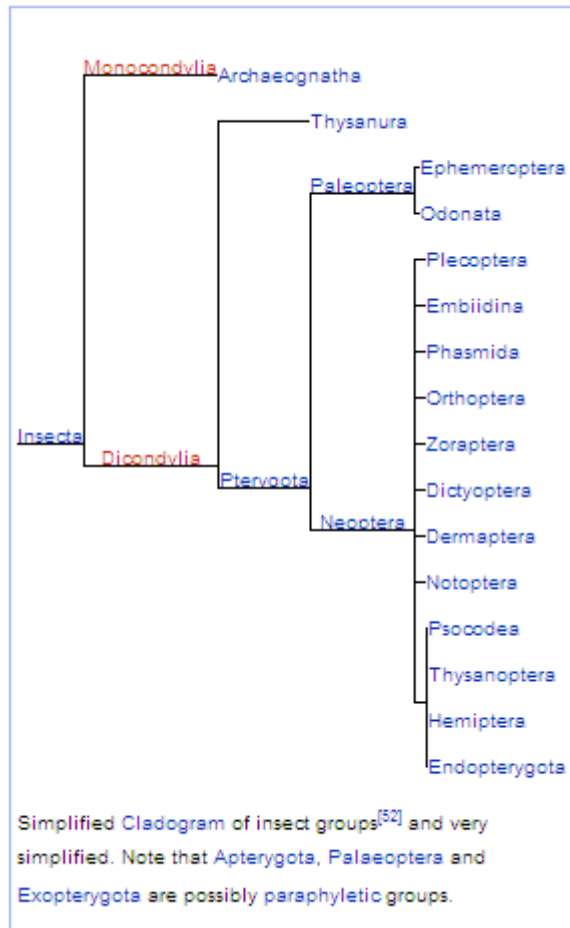


- Hymenoptera (ants, bees, etc.)
- Coleoptera (beetles)
- Strepsiptera (twisted-winged parasites)



not

- ~~Hymenoptera (ants, bees, etc.)~~
- Coleoptera (beetles)
- Strepsiptera (twisted-winged parasites)



- Hymenoptera (ants, bees, etc.)
- Coleoptera (beetles)
- Strepsiptera (twisted-winged parasites)



Beetle-Dermatitis

Paederus-Species

Cantharidin, Paederin





Ladybug

*Harmonia
axyridis*



*Larvae of
Harmonia
axyridis*

Asian ladybugs (*Harmonia axyridis*): A new seasonal indoor allergen

Takuya Nakazawa, MD, PhD,^a Shama M. Satinover, MS,^a Lisa Naccara, BA,^a
Lucy Goddard, RN,^a Bojan P. Dragulev, PhD,^b Edward Peters, MD,^c and
Thomas A. E. Platts-Mills, MD, PhD^a Charlottesville, Va, and Austin, Tex

TABLE I. Twenty patients reporting symptoms to ALB: environmental exposure and serum IgE antibody levels

	Age (y)/race/ sex	Symptoms	Season	Skin test	Household infestation	ALB	IgE antibody (IU/mL)			
							CR	Mite	Cat	Other*
1	46/W/F	Rhinitis, cough	Winter	ND	+++	59	1.5	2.1	Neg	Moth
2	60/W/M	Cough, asthma	Year round	++	++	30	5	10.6	10.6	Grass, moth, Berlin beetle
3	54/W/M	Rhinitis, cough	Winter	++	+++	41	Neg	Neg	Neg	Grass
4	56/W/F	Rhinitis, cough	Winter	ND	++	14	2.3	Neg	Neg	Grass, Berlin beetle
5	57/W/F	Asthma, rhinitis	Winter	++	++	10	0.4	22	8.9	Dog, moth, grass
6	72/W/F	Asthma, rhinitis	Winter	++	++	1.6	0.6	0.4	Neg	Moth†
7	58/W/F	Cough, conjunctivitis	Winter	++	++	4.1	Neg	Neg	Neg	Neg
8	18/W/M	Rhinitis	Spring	ND	+	Neg	1.5	Neg	Neg	Moth‡
9	47/W/F	Asthma	Fall	++	+++	26	Neg	Neg	Neg	Neg
10	59/W/F	Asthma	Year round	+	++	1.6	2.3	1.8	3.1	Dog, moth, grass, Berlin beetle
11	78/W/F	Asthma	Worse in winter	++	++	0.6	Neg	Neg	Neg	Neg
12	42/W/F	Rhinitis	Spring/fall	ND	+	Neg	Neg	1.2	Neg	Neg
13†	51/W/F	Rhinitis, conjunctivitis	Fall	ND	++	14.5	Neg	Neg	1.0	Neg
14	59/W/F	Chronic cough, rhinitis	Year round	ND	++	0.38	Neg	Neg	Neg	Neg
15	45/W/F	Asthma, rhinitis	Year round	ND	+	0.4	Neg	Neg	Neg	Grass
16	45/W/F	Asthma, hives	Fall	ND	++	Neg	Neg	Neg	Neg	Moth
17	63/W/F	Rhinitis, cough	Winter	ND	+++	3.6	Neg	Neg	Neg	Neg
18	49/W/F	Asthma	Winter	++	++	Neg	Neg	Neg	Neg	Neg
19	39/W/F	Asthma	Year round	ND	+	Neg	Neg	15.2	1.1	Dog, moth
20	34/W/F	Asthma	Year round	++	+++	0.68	0.67	19.6	0.7	Grass, moth

CR, Cockroach; F, female; M, male; W, white.

*Other allergens tested included rye grass, moth, Berlin beetle, and dog.

†Patient #13 reported a large local swelling of the neck after being bitten by a ladybug.

‡The IgE antibody responses to moth included 2 sera with high titer.

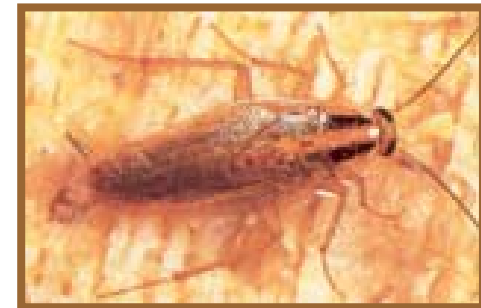
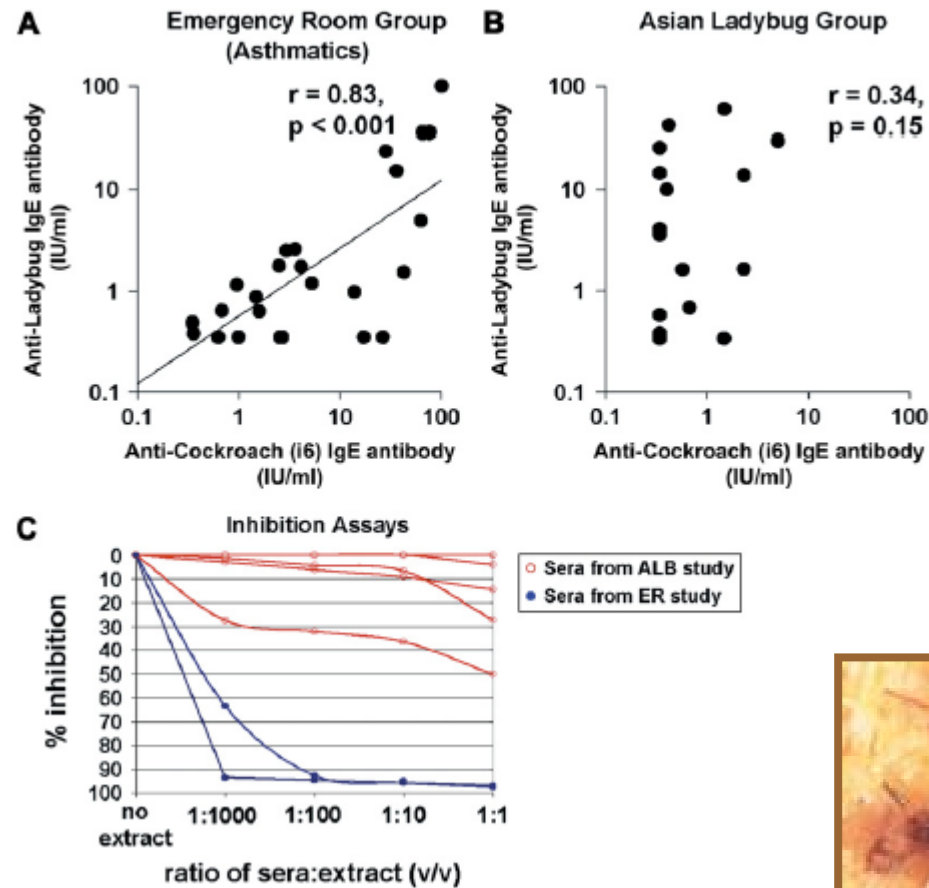
Ladybug - Allergy

History	Work with wheat, Seasonal RCA (winter)
Skin test	Extracts (Not com avail)
IgE	ImmunoCAP
Therapy	Symptomatically Eradication (SIT)



Crossreactivity

Ladybugs and Cockroaches



Cockroaches

Cockroaches



German cockroach



Brown-banded cockroach



American cockroach



Australian cockroach



Smoky-brown cockroach



Brown cockroach



Florida woods cockroach



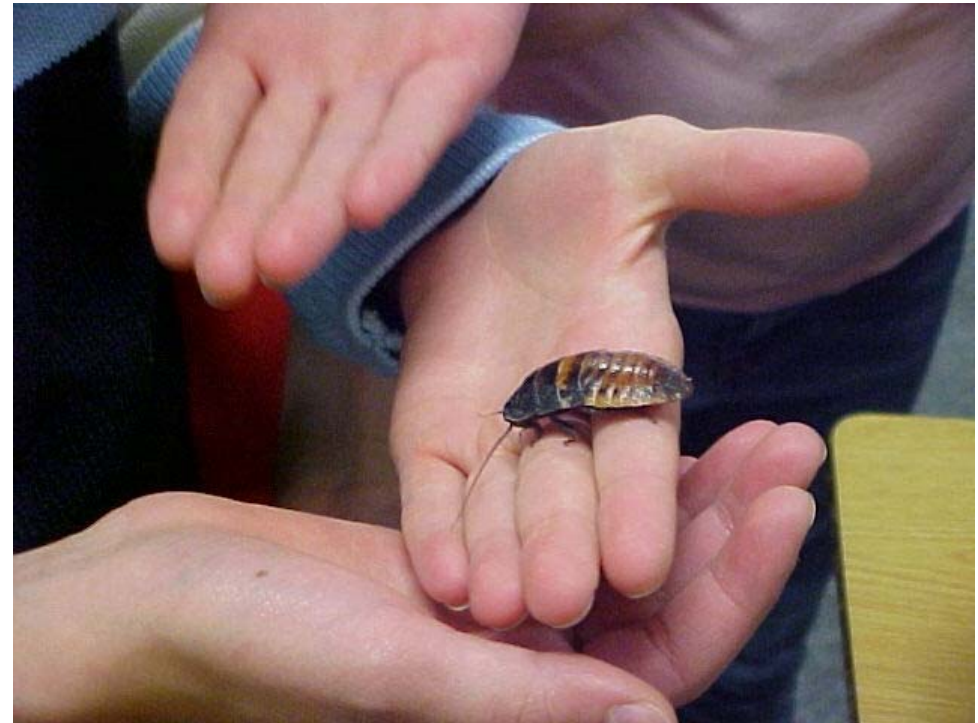
Surinam cockroach



Oriental cockroach



Cuban cockroach



Cockroaches

Current reviews of allergy and clinical immunology

(Supported by a grant from Astra Pharmaceuticals, Westborough, Mass)

Series editor: Harold S. Nelson, MD

Cockroach allergens and asthma

L. Karla Arruda, MD, PhD,^{a,b} Lisa D. Vailes, MS,^c Virginia P. L. Ferriani, MD, PhD,^a
Ana Beatriz R. Santos, BSc,^b Anna Pomés, PhD,^c and Martin D. Chapman, PhD^c
Ribeirão Preto, Brazil, and Charlottesville, Va

Although cockroach allergens are found throughout the house, including beds, furniture, and carpets, the highest levels are typically found in the kitchen, and these levels are perhaps the best indicator of cockroach infestation in a house.^{8,14,19} However, exposure in the bedroom and family room may be more relevant in causing sensitization.

J Allergy Clin Immunol 2005

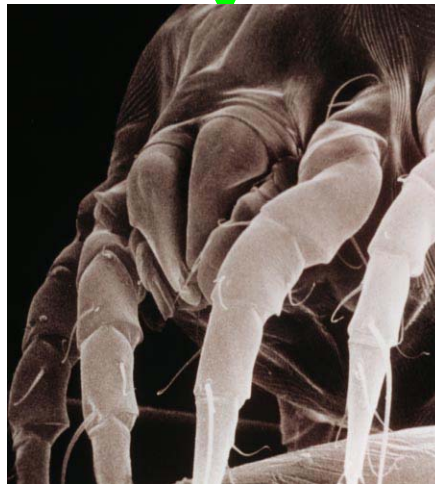




Seafood

Tropomyosin

Pen a 1 u.a.



Housedust-Mite



Cockroaches

Helbling A. Schw Med Wschr 1997
Santos AB, J Allergy Clin Immunol 1997
Kutting B, Brehler R. Hautarzt 2001

Mite-Crustazeae-Syndrome

What's on an allergologist's mind....



Tropomyosin
Pen a 1



Housedust-Mite
Cockroach



Seafood

Red Mosquito - Mite-Crustazeae-Syndrome

What's on an allergologist's mind....

Tropomyosin

Pen a 1

Housedust-Mite

Seafood



Mosquito allergy



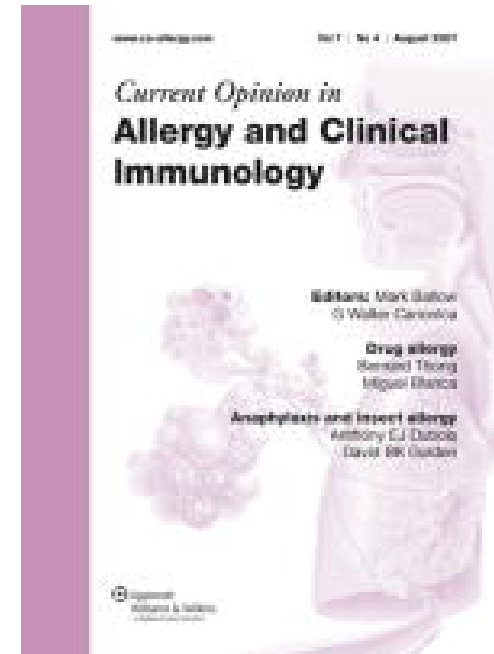
Local reaction

f (Host)

f (Mosquito)



You don't feel the sting
Only the reaction



Advances in mosquito allergy.

Peng, Zhikang; Simons, F
Current Opinion in Allergy & Clinical Immunology. 7(4):350-354, August 2007.

Mosquito allergy:

Allergens are in the saliva

Table 1 *Aedes aegypti* salivary proteins

Protein name	Allergen name	Molecular weight (kDa)	cDNA sequenced	Biological functions
α -Amylase 1		81.5	Yes	Unknown
apyrase	Aed a 1	68	Yes	Antiplatelet
α -Glucosidase (maltase-like 1)	Aed a 4	67	Yes	Sugar digestion
Esterase		65		Unknown
Anticoagulant-factor Xa		54	Yes	Anticoagulant
Aed a X ₁	Aed a X ₁ ?	44		Unknown
Aed a X ₂	Aed a X ₂ ?	37		Unknown
Female-specific protein, D7	Aed a 2	37	Yes	Unknown
	Aed a 3	30	Yes	Unknown
Sialokinins		1.4	Yes	Vasodilator
Antitumour necrosis factor		Unknown		Antitumour
Lysozyme		Unknown		Bacteriolysis

Reproduced from Peng and Simons [1].

Peng, Zhikang; Simons, F
Current Opinion in Allergy & Clinical Immunology. 7(4):350-354, August 2007.

Mosquito Allergy

History

Mostly large local reactions
Systemic allergic symptoms < 3%

Skin test

Extracts (Com. Available, Sens +/-)
Recombinant allergens

IgE

ELISA >ImmunoCAP

Therapy

Symptomatically
Prevention (Repellents, Long sleeves
SIT with WBA



Mosquitoes as sources of inhalant allergens: Clinicoimmunologic and biochemical studies

236 patients in North-India with Asthma

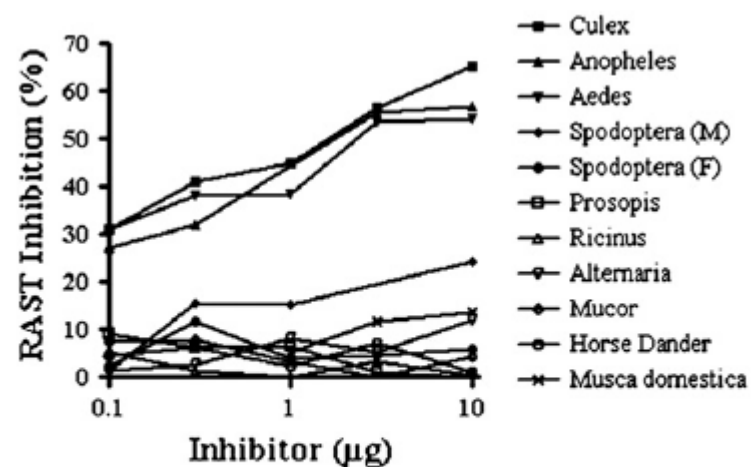


FIG 1. Inhibition of mosquito (*C quinquefasciatus*) RAST results with homologous (*C quinquefasciatus*) and the 2 heterologous mosquito WBEs (*A aegypti* and *A stephensi*) and 8 unrelated allergen extracts.







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Rohkalium 0,10%
Rohmagnesium 0,10%
Rohzink 0,10%
Rohkupfer 0,10%
Rohmangan 0,10%
Rohnatrium 0,10%
Rohchlorid 0,10%
Rohsilber 0,10%
Rohblei 0,10%
Rohzinn 0,10%
Rohzink 0,10%
Rohkupfer 0,10%
Rohmangan 0,10%
Rohnatrium 0,10%
Rohchlorid 0,10%
Rohsilber 0,10%
Rohblei 0,10%
Rohzinn 0,10%



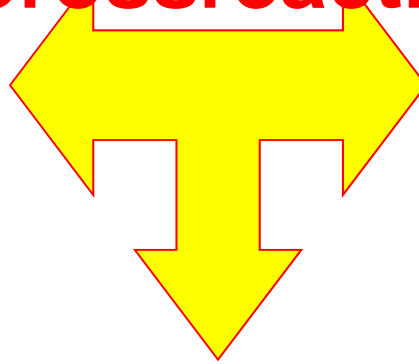
Red mosquito larvae

Allergy against Red mosquito larvae

History	Feeding fishes, Preparing fish food
Skin test	Extracts (Not com avail)
IgE	ImmunoCAP
Therapy	Symptomatically Change of fish food



Crossreactivity



Tropomyosin

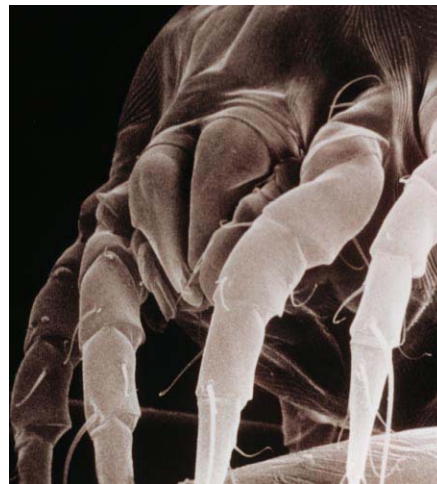
Pen a 1 u.a.

Red Mosquito larvae

Chironomids



Cockroaches



Housedust-Mite

Johansson E Allergy 2001

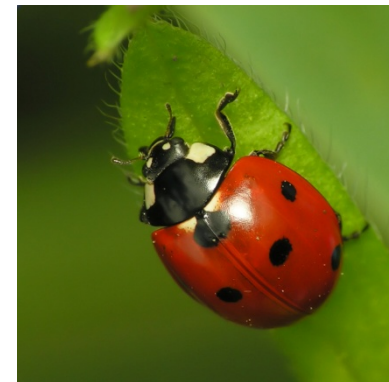
Insect bites and stings

Most common allergens



photo (c) Alex Wild

iRFA





Danke schön

Merci

Gracias

Thank you

Grazie

peter.schmid@usz.ch